Applicant: George E. Wright Attorney Docket No.: 07917-0183001 / UMMC 03-23

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AMENDMENTS TO THE CLAIMS:

This listing of claims replaces all prior versions and listings of claims in the application:

Listing of Claims:

Please cancel claims 4, 6, 9, and 35, and amend the claims as indicated below.

- 1. (Currently Amended) A composition comprising a combination of:
- a) an inhibitor of Herpes simplex virus thymidine kinase that is selected from the group consisting of 2-phenylamino-9-substituted-6-oxopurines and 2-phenylamino-9H-6-oxopurines, or an ester, salt, or solvate thereof, in a first dose less than a median therapeutically effective dose of the inhibitor of Herpes simplex virus thymidine kinase, and
- b) an antiherpes substance that inhibits viral DNA replication comprising one or more of (1) a pre-phosphorylated or phosphonate nucleoside analog; (2) a pyrophosphate analog; and (3) a nucleoside analog, or any combination thereof, or an ester, salt, or solvate thereof, in a second dose less than a median therapeutically effective dose of the antiherpes substance;
- wherein the first dose and the second dose together form a therapeutically effective dose of the combination.
- (Original) The composition of claim 1, wherein the antiherpes substance comprises one or more of (1) a pre-phosphorylated or phosphonate nucleoside analog or (2) a pyrophosphate analog, or any combination thereof, or an ester, salt, or solvate thereof.
- 3. (Original) The composition of claim 1, wherein the pre-phosphorylated or phosphonate nucleoside analog is acyclovir monophosphate, ganciclovir monophosphate, cidofovir, or 9-(phosphonomethoxyethyl)adenine (PMEA), or an ester, salt, or solvate thereof.
- 4. (Cancelled).
- (Original) The composition of claim 1, wherein the nucleoside analog is acyclovir, famiciclovir, or ganciclovir, or an ester, salt, or solvate thereof.
- (Cancelled).

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7. (Previously Presented) The composition of claim 1, wherein the inhibitor of Herpes simplex virus thymidine kinase is selected from the group consisting of 2-phenylamino-9-substituted-6-

oxopurines, or an ester, salt, or solvate thereof.

8. (Original) The composition of claim 1, wherein the thymidine kinase inhibitor is 2-

phenylamino-9-(4-hydroxybutyl)-6-oxopurine, or an ester, salt, or solvate thereof.

9. (Cancelled).

10. (Original) The composition of claim 1, wherein the antiherpes substance is cidofovir or an

ester or salt thereof.

11. (Original) The composition of claim 1, wherein the antiherpes substance is acyclovir, or an

ester, salt, or solvate thereof.

12. (Original) The composition of claim 1, wherein the antiherpes substance is acyclovir

monophosphate, or an ester, salt, or solvate thereof.

13. (Original) The composition of claim 1, wherein the antiherpes substance is ganciclovir

monophosphate, or an ester, salt, or solvate thereof.

 $14. \ \, (Original) \ \, A \ \, dosage \ \, form \ \, for \ \, parenteral \ \, or \ \, oral \ \, use \ \, containing \ \, a \ \, pharmaceutical$

composition according to claim 1.

15. (Original) A cream, lotion, gel, ointment, plaster, stick, or pen containing a

composition according to claim 1.

16. (Original) The composition of claim 1, including a pharmaceutically acceptable carrier that

is selected from the group consisting of sterile water, saline, polyalkylene glycols, vegetable oils.

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hydrogenated naphthalenes, biocompatible polymers, biodegradable polymers, and mixtures thereof.

17. (Original) The composition of claim 16, wherein the biodegradable polymer is selected from the group consisting of polycaprolactone, polydecalactone, poly(sebacic anhydride). sebacic acid-co-1,3-bis(carboxyphenoxypropane), sebacic acid-co-1,6-

bis(carboxypheno 1,6-bis(carboxyphenoxyhexane), albumin and derivatives, gelatin and derivatives, starch and derivatives, gum arabic, cellulose and derivatives, polysorbate and derivatives, agarose, lectins, galactose, polyurethanes, polyvinylalcohol, functionalized polymers and copolymers of lactic and glycolic acid, lactic acid homopolymer, glycolic acid copolymer, copolymers of lactic acid and glycolic acid, polyhydroxybutyrate, polyhydroxyalkanoic acid, and mixtures thereof.

- 18. (Original) The composition of claim 17, wherein the biodegradable polymer is in the form of a particle.
- 19. (Original) The composition of claim 18, wherein the particle includes multiple walls.

20-31. (Cancelled)

- 32. (Currently Amended) A kit for treatment of a Herpes simplex virus infection in a mammal, the kit comprising:
- a) an inhibitor of Herpes simplex virus thymidine kinase that is selected from the group consisting of 2-phenylamino-9-substituted-6-oxopurines and 2-phenylamino-9H-6-oxopurines, or an ester, salt, or solvate thereof, in a first dose less than a median therapeutically effective dose of the inhibitor of Herpes simplex virus thymidine kinase.
- b) an antiherpes substance that inhibits viral DNA replication comprising one or more of (1) a pre-phosphorylated or phosphonate nucleoside analog; (2) a pyrophosphate analog; and (3) a nucleoside analog, or any combination thereof, or an ester, salt or solvate thereof, in a second dose less than a median therapeutically effective dose of the antiherpes substance:

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wherein the first dose and the second dose together form a therapeutically effective dose of the combination; and

c) instructions for administering (a) and (b) concurrently or within a sufficiently close time to achieve coexistent concentrations of (a) and (b) in subject.

- 33. (Original) The kit of claim 32, wherein the pre-phosphorylated or phosphonate nucleoside analog is acyclovir monophosphate, ganciclovir monophosphate, cidofovir, or 9-(phosphonomethoxyethyl)adenine (PMEA), or an ester, salt, or solvate thereof.
- 34. (Previously Presented) The kit of claim 32, wherein the inhibitor of Herpes simplex virus thymidine kinase is selected from the group consisting of 2-phenylamino-9-substituted-6oxopurines, or an ester, salt, or solvate thereof.
- 35. (Cancelled).
- 36. (Original) The kit of claim 32, wherein the antiherpes substance is cidofovir or an ester or salt thereof.
- 37. (Previously Presented) The kit of claim 32, wherein the antiherpes substance is acyclovir, or an ester, salt, or solvate thereof.
- 38. (Previously Presented) The kit of claim 32, wherein the antiherpes substance is acyclovir monophosphate, or an ester, salt, or solvate thereof.
- 39. (Previously Presented) The kit of claim 32, wherein the thymidine kinase inhibitor is 2phenylamino-9-(4-hydroxybutyl)-6-oxopurine, or an ester, salt, or solvate thereof, and wherein the antiherpes substance is acyclovir, or an ester, salt, or solvate thereof.

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40. (Previously Presented) The composition of claim 1, wherein the thymidine kinase inhibitor is 2-phenylamino-9-(4-hydroxybutyl)-6-oxopurine, or an ester, salt, or solvate thereof, and

wherein the antiherpes substance is acyclovir, or an ester, salt, or solvate thereof.

41. (Withdrawn) A method for treating a recurrent Herpes simplex virus infection in a mammal, the method comprising administering to the mammal a therapeutic dose of the composition of

claim 1.

42. (Withdrawn) A method for treating a Herpes simplex virus infection in a mammal, the

method comprising obtaining the kit of claim 32 and administering the inhibitor and antiherpes

substance according to the instructions.

43. (Currently Amended) A composition comprising a combination of:

a) <u>a 2-phenylamino-9-(4-hydroxybutyl)-6-oxopurine</u>, or an ester, salt, or solvate thereof

in a first dose less than a median therapeutically effective dose; and

b) an antiherpes substance selected from the group consisting of foscarnet or an ester,

salt, or solvate thereof, acyclovir, or an ester, salt, or solvate thereof, and cidofovir or an ester, salt, or solvate thereof, in a second dose less than a median theraneutically effective dose:

wherein the first dose and the second dose together form a therapeutically effective dose

of the combination.

44. (Previously Presented) The composition of claim 43, wherein the antiherpes substance

comprises acyclovir or an ester, salt, or solvate thereof.

45. (Currently Amended) A kit for treatment of a Herpes simplex virus infection in a mammal,

the kit comprising:

a) a 2-phenylamino-9-(4-hydroxybutyl)-6-oxopurine, or an ester, salt, or solvate thereof

in a first dose less than a median therapeutically effective dose;

b) an antiherpes substance selected from the group consisting of foscarnet or an ester,

salt, or solvate thereof, acyclovir, or an ester, salt, or solvate thereof, and cidofovir or an ester,

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salt, or solvate thereof, in a second dose less than a median therapeutically effective dose. wherein the first dose and the second dose together form a therapeutically effective dose; and

c) instructions for administering (a) and (b) concurrently or within a sufficiently close time to achieve coexistent concentrations of (a) and (b) in subject.

46. (Previously Presented) The kit of claim 45, wherein the antiherpes substance comprises acyclovir or an ester, salt, or solvate thereof.

Please enter the following new claims.

47. (New) The composition of claim 8, wherein the nucleoside analog is selected from the group consisting of acyclovir, famiciclovir, or ganciclovir, and an ester, salt, or solvate thereof.

48. (New) The composition of claim 8, wherein the antiherpes substance is selected from the group consisting of acyclovir, acyclovir monophosphate, and an ester, salt, or solvate thereof.

49. (New) The composition of claim 1, wherein

- a) the inhibitor of Herpes simplex virus thymidine kinase is 2-phenylamino-9-(4hydroxybutyl)-6-oxopurine, or an ester, salt, or solvate thereof; and
- the antiherpes substance that inhibits viral DNA replication is selected from the group consisting of acyclovir, famiciclovir, ganciclovir, and an ester, salt, or solvate thereof.
- 50. (New) The composition of claim 1, wherein the subject is a mammal.